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NEWS	3	APR 28	The DWPI (files WPINDEX, WPIDS and WPIX) on STN have been enhanced with thesauri for the European Patent Classifications
NEWS	4	MAY 02	MEDLINE Improvements Provide Fast and Simple Access to DOI and Chemical Name Information
NEWS	5	MAY 12	European Patent Classification thesauri added to the INPADOC files, PCTFULL, GBFULL and FRFULL
NEWS	6	MAY 23	Enhanced performance of STN biosequence searches
NEWS	7	JUN 20	STN on the Web Enhanced with New Patent Family Assistant and Updated Structure Plug-In
NEWS	8	JUN 20	INPADOC databases enhanced with first page images
NEWS	9	JUN 20	PATDPA database updates to end in June 2011
NEWS	10	JUN 26	MARPAT Enhancements Save Time and Increase Usability
NEWS	11	JUL 25	STN adds Australian patent full-text database, AUPATFULL, including the new numeric search feature.
NEWS	12	AUG 01	CA Sections Added to ACS Publications Web Editions Platform
NEWS	13	AUG 16	INPADOC: Coverage of German Patent Data resumed, enhanced legal status
NEWS	14	AUG 18	Upgrade now to STN Express, Version 8.5
NEWS	15	SEP 01	CAS Journal Coverage Now Includes Ahead-of-Print Articles for More Than 100 Journal Titles
NEWS	16	SEP 01	Older Versions of STN Express to be Discontinued Beginning in March 2012
NEWS	17	SEP 09	USAN Database Updates Offer Superior Currency on STN(R)
NEWS	18	SEP 26	STN Adds Canadian Patent Full-text Database - CANPATFULL
NEWS	19	SEP 26	GEOREF and ENCOMPLIT databases were reloaded on September 24, 2011.
NEWS	20	SEP 26	Updates to the IFIPAT/IFIUDB/IFICDB databases have resumed.
NEWS	21	SEP 26	ECLA Thesaurus in CA/CAPLUS Improves Patent Searching on STN
NEWS	22	SEP 26	Access AUPATFULL and CANPATFULL databases with STN Viewer
NEWS	23	OCT 26	New STN Revolutionizes Patent Searching for Professionals
NEWS	24	DEC 1	CA/CAPLUS Now Includes Examiner Citations for Japanese Patents
NEWS	25	DEC 1	CAS Expands Global Patent Coverage - Intellectual Property Corporation of Malaysia Becomes 62nd Authority on CA/CAPLUS
NEWS	26	DEC 5	STN on the Web Enhancements Include Compatibility with Microsoft Windows 7
NEWS	27	DEC 14	Removal of ITRD and PATIPC databases from STN
NEWS	28	DEC 15	Rolled-up IPC Core Codes Removed from IPC Reclassifications in Patent Databases on STN
NEWS EXPRESS	18	AUGUST 2011	CURRENT WINDOWS VERSION IS V8.5, AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2011.

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 16:16:01 ON 04 JAN 2012

```
=> fil medline biosis caplus scisearch embase wpids
COST IN U.S. DOLLARS          SINCE FILE      TOTAL
                              ENTRY      SESSION
FULL ESTIMATED COST          0.24      0.24
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FILE 'MEDLINE' ENTERED AT 16:16:39 ON 04 JAN 2012

FILE 'BIOSIS' ENTERED AT 16:16:39 ON 04 JAN 2012
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FILE 'WPIDS' ENTERED AT 16:16:39 ON 04 JAN 2012
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```
=> s hairpin(L)ligase
L1      833 HAIRPIN(L) LIGASE

=> s hairpin(L)ligase(L)(adapter or connector)
L2      17 HAIRPIN(L) LIGASE(L) (ADAPTER OR CONNECTOR)
```

```
=> dup rem
ENTER L# LIST OR (END):12
PROCESSING COMPLETED FOR L2
L3      13 DUP REM L2 (4 DUPLICATES REMOVED)
```

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=> d ibib abs 1
```

```
L3  ANSWER 1 OF 13      MEDLINE ® on STN      DUPLICATE 1
ACCESSION NUMBER: 2011850744      MEDLINE
DOCUMENT NUMBER: PubMed ID: 21876038
TITLE: TLR regulation of SPSB1 controls inducible nitric oxide
synthase induction.
AUTHOR: Lewis Rowena S; Kolesnik Tatiana B; Kuang Zhihe; D'Cruz
```

CORPORATE SOURCE: Akshay A; Blewitt Marnie E; Masters Seth L; Low Andrew;
 Willson Tracy; Norton Raymond S; Nicholson Sandra E
 Walter and Eliza Hall Institute of Medical Research,
 Parkville, Victoria 3052, Australia.
 SOURCE: Journal of immunology (Baltimore, Md. : 1950), (2011 Oct 1)
 Vol. 187, No. 7, pp. 3798-805. Electronic Publication:
 2011-08-29.
 Journal code: 2985117R. E-ISSN: 1550-6606. L-ISSN:
 0022-1767.
 PUB. COUNTRY: United States
 DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
 (RESEARCH SUPPORT, NON-U.S. GOV'T)
 LANGUAGE: English
 FILE SEGMENT: Abridged Index Medicus Journals; Priority Journals
 ENTRY MONTH: 201111
 ENTRY DATE: Entered STN: 22 Sep 2011
 Last Updated on STN: 16 Dec 2011
 Entered Medline: 22 Nov 2011

AB The mammalian innate immune system has evolved to recognize foreign
 molecules derived from pathogens via the TLRs. TLR3 and TLR4 can signal
 via the TIR domain-containing adapter inducing IFN- β (TRIF), which
 results in the transcription of a small array of genes, including
 IFN- β . Inducible NO synthase (iNOS), which catalyzes the production
 of NO, is induced by a range of stimuli, including cytokines and microbes.
 NO is a potent source of reactive nitrogen species that play an important
 role in killing intracellular pathogens and forms a crucial component of
 host defense. We have recently identified iNOS as a target of the
 mammalian SPSB2 protein. The SOCS box is a peptide motif, which, in
 conjunction with elongins B and C, recruits cullin-5 and Rbx-2 to form an
 active E3 ubiquitin ligase complex. In this study, we show that SPSB1
 is the only SPSB family member to be regulated by the same TLR pathways
 that induce iNOS expression and characterize the interaction between SPSB1
 and iNOS. Through the use of SPSB1 transgenic mouse macrophages and short
 hairpin RNA knockdown of SPSB1, we show that SPSB1 controls both the
 induction of iNOS and the subsequent production of NO downstream of TLR3
 and TLR4. Further, we demonstrate that regulation of iNOS by SPSB1 is
 dependent on the proteasome. These results suggest that SPSB1 acts
 through a negative-feedback loop that, together with SPSB2, controls the
 extent of iNOS induction and NO production.

=> d his

(FILE 'HOME' ENTERED AT 16:16:01 ON 04 JAN 2012)

FILE 'MEDLINE, BIOSIS, CAPLUS, SCISEARCH, EMBASE, WPIDS' ENTERED AT
 16:16:39 ON 04 JAN 2012

L1 833 S HAIRPIN(L)LIGASE
 L2 17 S HAIRPIN(L)LIGASE(L) (ADAPTER OR CONNECTOR)
 L3 13 DUP REM L2 (4 DUPLICATES REMOVED)

=> s l3 and py<=2003
 L4 6 L3 AND PY<=2003

=> d ibib abs 1-6

L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2012 ACS on STN
 ACCESSION NUMBER: 1992:36605 CAPLUS
 DOCUMENT NUMBER: 116:36605
 ORIGINAL REFERENCE NO.: 116:6161a,6164a
 TITLE: Modifying specificities of restriction enzymes
 AUTHOR(S): Szybalski, Wacław

CORPORATE SOURCE: McArdle Lab. Cancer Res., Univ. Wisconsin, Madison, WI, 53706, USA

SOURCE: Biotechnol.: Bridging Res. Appl., Proc. U.S.-Isr. Res. Conf. Adv. Appl. Biotechnol. (1991), Meeting Date 1990, 371-6. Editor(s): Kamely, Daphne; Chakrabarty, Ananda M.; Kornguth, Steven E. Kluwer: Boston, Mass.
CODEN: 57MWA2

DOCUMENT TYPE: Conference; General Review

LANGUAGE: English

AB A review with 12 refs. Using restriction enzymes, modification methyltransferases, single-strand-specific nucleases, ligases, oligo hairpin adapters, and DNA-binding proteins, the authors have designed various strategies for very specific cleavage of small, medium, and very large DNA mols., for detecting point mutations or other genetic changes, for a very efficient printing reaction to produce tetranucleotides, as a highly amplified detection signal, or to synthesize copious amts. of longer sequences, for development of trimming, gene-fusion, or amplification plasmids, and for various other applications, some already tested and some in various stages of development.

L4 ANSWER 2 OF 6 WPIDS COPYRIGHT 2012 THOMSON REUTERS on STN

ACCESSION NUMBER: 2003-021218 [200302] WPIDS

DOC. NO. CPI: C2003-005375 [200302]

TITLE: Selectively amplifying unknown DNA sequence, useful when analyzing single nucleotide polymorphism, by digesting DNA into fragments with single-strand cohesive ends, ligating fragments with a hairpin loop adapter and amplifying the fragments

DERWENT CLASS: B04; D16

INVENTOR: JEON J; JEON J T; JOUNG I; JUN J T; JUNG I S; LEE J W; PARK H; PARK H O; RHEE J; SONG S; SONG S N; WEON S; WEON S Y; WON S Y; JOUNG I S; RHEE J W

PATENT ASSIGNEE: (BION-N) BIONEER CORP; (BION-N) BIONIA JH; (JEON-I) JEON J; (JOUN-I) JOUNG I; (PARK-I) PARK H O; (RHEE-I) RHEE J; (SONG-I) SONG S; (WEON-I) WEON S

COUNTRY COUNT: 30

PATENT INFO ABBR.:

PATENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC
EP 1256630	A2	20021113	(200302)*	EN	10[4]	<--
US 20020192769	A1	20021219	(200315)#	EN		<--
CA 2344599	A1	20021107	(200316)#	EN		<--
JP 2003009864	A	20030114	(200316)	JA	6	<--
KR 2002085727	A	20021116	(200320)	KO		<--
US 6849404	B2	20050201	(200511)#	EN		
KR 758610	B1	20070913	(200839)	KO		
EP 1256630	B1	20080618	(200841)	EN		
DE 60227130	E	20080731	(200853)	DE		
JP 4435463	B2	20100317	(201020)	JA	8	
CA 2344599	C	20110712	(201149)#	EN		

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE

EP 1256630 A2
 CA 2344599 A1
 KR 2002085727 A
 KR 758610 B1
 US 20020192769 A1
 US 6849404 B2
 DE 60227130 E
 DE 60227130 E
 JP 2003009864 A
 JP 4435463 B2
 CA 2344599 C

EP 2002-10053 20020506
 CA 2001-2344599 20010507
 KR 2001-25637 20010507
 KR 2001-25637 20010507
 US 2001-849597 20010507
 US 2001-849597 20010507
 DE 2002-60227130 20020506
 EP 2002-10053 20020506
 JP 2002-131307 20020507
 JP 2002-131307 20020507
 CA 2001-2344599 20010507

FILING DETAILS:

PATENT NO	KIND	PATENT NO
DE 60227130 E	Based on	EP 1256630 A
KR 758610 B1	Previous Publ	KR 2002085727 A
JP 4435463 B2	Previous Publ	JP 2003009864 A

PRIORITY APPLN. INFO: KR 2001-25637 20010507
 US 2001-849597 20010507
 CA 2001-2344599 20010507

AN 2003-021218 [200302] WPIDS

AB EP 1256630 A2 UPAB: 20100323

NOVELTY - A process for selective amplifying DNA of which base sequence is completely unknown, comprising digesting DNA into fragments having a single-strand cohesive end group, ligating the DNA fragments with a hairpin loop adaptor having a single-strand cohesive end which can be complementarily combined and ligated on both ends of the DNA, and amplifying the fragments using DNA polymerase and primer.

DETAILED DESCRIPTION - A process (M1) for selective amplifying DNA of which base sequence is completely unknown, comprising:

(a) a step for digesting DNA into fragments which has a single-strand cohesive end group by using restriction enzyme, and separately from the above step, a step for preparing hairpin loop adaptor which has the single-strand cohesive end which can be complementarily combined and ligated on the both ends of the DNA fragments obtained in the above;

(b) a step for ligating the DNA fragments with the hairpin loop adaptor thus prepared by using DNA ligase;

(c) a step for removing DNA fragments and hairpin loop adaptor which have not participated in the ligation reaction by using exonuclease; and

(d) a step for amplifying the DNA fragment by using DNA polymerase and primer which can combine complementarily on the residual sequence from the adaptor.

INDEPENDENT CLAIMS are included for the following:

(1) a process for making library of DNA fragment of which terminal sequence are known by using DNA of which base sequence is completely unknown, comprising:

(a) a step for digesting DNA into fragments which have single-strand cohesive end by using restriction enzyme, and separately from the above, for preparing a series of hairpin loop adapters which have single-strand cohesive end of which base sequence is known;

(b) a step for ligating the DNA fragments with the hairpin loop adapters prepared in the above step (a) by using DNA ligase; and

(c) a step for eliminating the hairpin loop only from the DNA fragments which contain hairpin loop adapter, obtained in step (b) by treating alkaline solution, RNase of single strand specific exonuclease; and

(2) a series of hairpin loop adapters which have single-strand

cohesive ends, where the single-strand cohesive ends are composed of all sorts of single-strand DNA which can be made by random combination of four nucleotides.

USE - The process is useful for analyzing single nucleotide polymorphism in the nucleotide sequences of each individual.

L4 ANSWER 3 OF 6 WPIDS COPYRIGHT 2012 THOMSON REUTERS on STN
 ACCESSION NUMBER: 2002-617299 [200266] WPIDS
 CROSS REFERENCE: 2001-040475; 2001-366407; 2002-048724
 DOC. NO. CPI: C2002-174481 [200266]
 TITLE: Isolating DNA containing fragments nicked by Escherichia coli methyl-directed mismatch repair system employs a modified rolling circle amplification procedure which utilizes DNA polymerase III
 DERWENT CLASS: B04; D16
 INVENTOR: LASKEN R; WEISSMAN S
 PATENT ASSIGNEE: (LASK-I) LASKEN R; (MOLE-N) MOLECULAR STAGING INC; (UYYA-C) UNIV YALE; (WEIS-I) WEISSMAN S
 COUNTRY COUNT: 1

PATENT INFO ABBR.:

PATENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC	
US 20020076704	A1	20020620	(200266)*	EN	19[5]		<--
US 6576448	B2	20030610	(200340)	EN			<--

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 20020076704	A1	Provisional	US 1998-100996P 19980918
US 20020076704	A1	Div Ex	US 1999-398216 19990917
US 20020076704	A1		US 2001-818927 20010328
US 6576448	B2	Provisional	US 1998-100996P 19980918
US 6576448	B2	Div Ex	US 1999-398216 19990917
US 6576448	B2		US 2001-818927 20010328

FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 6576448	B2	US 6235502
	Div ex	B

PRIORITY APPLN. INFO: US 2001-818927 20010328
 US 1998-100996P 19980918
 US 1999-398216 19990917

AN 2002-617299 [200266] WPIDS
 CR 2001-040475; 2001-366407; 2002-048724
 AB US 20020076704 A1 UPAB: 20060120

NOVELTY - Mismatched DNA containing fragments nicked by MutSLH from two DNA samples are isolated by a modified rolling circle amplification (RCA) method. DNA sample is digested with restriction enzyme, the fragments ligated to Y-shaped adapters forming adapter-fragment constructs which are subjected to nicking by MutSLH proteins and nicked fragments are treated with DNA polymerase(s) to elongate the 3' end.

DETAILED DESCRIPTION - Isolating (M1) mismatched DNA containing fragments nicked by MutSLH (a combination of the three proteins of the Escherichia coli methyl-directed mismatch repair system, collectively called MutSLH) from two DNA samples, comprises:

(a) digesting a first DNA sample with a restriction enzyme to

obtain DNA fragments;

(b) ligating Y-shaped adapters to the fragments to obtain fragment-plus-adapter constructs;

(c) repeating the above steps with a second sample;.

(d) methylating the products formed from (c) of the first sample but not the second sample;

(e) mixing the methylated and unmethylated products obtained from the samples, denaturing and reannealing to form hemimethylated heterohybrids;

(f) treating the fragments produced in step (e) with a MutSLH preparation to nick DNA containing mismatches and form a 3'-OH end; and

(g) elongating the 3'-OH end with one or more DNA polymerases.

The method can also be carried out by ligating adapters to DNA fragments to obtain fragment-adapter constructs that have a single-stranded (ss) overhang on the 5'-ends and recessed 3'-ends, where the adapters are blocked with a dideoxynucleotide or other modification that prevents the elongation by a DNA polymerase, carrying out (c)-(e) as above, blocking pre-existing nicks on the fragments with dideoxynucleoside triphosphates or their analogs, treating the fragments with MutSLH preparation to form 3'-OH ends and elongating the 3'-OH ends with a DNA polymerase having strand displacement or nick translation capacity.

INDEPENDENT CLAIMS are also included for the following:

(1) amplifying (M2) DNA using RCA in a manner that results in approximately equimolar amplification of DNA, by amplifying a circle DNA molecule which comprises a stem-and-loop, or a circle DNA molecule containing a universal base, an abasic residue, or other residue that alters the rate of circle replication, where elongation of the circle by a replicating polymerase carrying out the amplification is inhibited;

(2) a method (M3) for using RCA to obtain approximately equimolar amplification of DNA fragment mixtures by introducing one pause site in a DNA circle, comprises:

(a) placing within the circle a stretch of one or more abasic sites, a region of high secondary structure and a ligand binding site, and carrying out RCA under conditions such that the pause site slows the DNA polymerase replicating the circle and tends to equalize the number of times the polymerase copies circles of different length; or

(b) circularizing double stranded DNA (dsDNA) fragments around an adapter such that one strand of the adapter has an internal pause and the outer strand is blocked from ligation at one or both ends; or

(c) using a splint that has an internal double stranded (ds) segment containing a pause site on one DNA strand, and which has single stranded extensions at both ends complementary to the ends of the single-stranded fragment to be amplified;

(3) a method (M4) in which fragments of DNA may be circularized and amplified, by cutting DNA with a restriction enzyme that produces a single fragment containing all the markers of the region having pause to be determined, or using the RecA-assisted restriction endonuclease (RARE) method or its variants to produce such a fragment, ligating the cut DNA around an adapter and amplifying the fragment using RCA to obtain a single stranded DNA (ssDNA) from the desired fragment;

(4) amplifying (M5) segments of a DNA target by RCA, by annealing a padlock DNA that creates a gap comprising the sequence to be amplified, extending the 3'-OH of the padlock DNA with a DNA polymerase which generates a nick, ligating the nick to form a circle DNA, annealing an RCA primer to the circle DNA and carrying out RCA with DNA polymerase III;

(5) a method (M6) in which large fragments of DNA may be circularized and amplified, by providing a DNA-adapter construct which has a DNA fragment ligated around an adapter that has an internal gap and 5' region of non-complementarity and amplifying the DNA-adapter construct using DNA pol III in a RCA;

(6) a method (M7) of ligating adapters onto ends of DNA fragments for use in RCA of the DNA, by ligating restriction endonuclease fragments

with an oligonucleotide adapter that forms a hairpin structure with itself such that 3' and 5' ends of the adapter are annealed together and form a proper overhang or blunt end for ligation to the restriction endonuclease fragments, where a circular construct is formed, annealing an oligonucleotide primer to the single-stranded loop portion of the adapters and extending the oligonucleotide primer with a DNA polymerase in RCA reaction; and

(7) a method of carrying out RCA reaction, where at least two different DNA polymerases are employed, one or more of which contains a 3' to 5' exonuclease activity capable of correcting nucleotide misincorporations.

USE - (M1) is useful for isolating mismatched DNA containing fragments nicked by MutSLH from two DNA samples. The procedure is useful for polymorphism analysis (claimed).

The procedure allows for the MutSLH nicking of DNA fragments containing mismatched bases to analyze polymorphism between DNA samples. The methods are useful in phase determination, polymorphism analyses, mismatch scanning procedures and cloning procedures, diagnostics, genotyping, genomic mapping, DNA sequencing and synthesis of DNA probes.

ADVANTAGE - DNA polymerase III holoenzyme provides a superior rate of DNA synthesis and also high processivity which allows rapid replication through regions of high GC content, hair-pin structures, and other regions of secondary structure and regions that normally slow replication due to local sequence context effects.

L4 ANSWER 4 OF 6 WPIDS COPYRIGHT 2012 THOMSON REUTERS on STN
 ACCESSION NUMBER: 2002-106604 [200214] WPIDS
 DOC. NO. CPI: C2002-032857 [200214]
 TITLE: Simultaneous sequence-specific identification and separation of polynucleotide fragments, comprises using restriction endonucleases that recognize degenerate bases in their recognition/cleavage sequence, useful in DNA fingerprinting
 DERWENT CLASS: B04; D16
 INVENTOR: LI B; LI B Y T M R I; SHI L; SHI L T M R I; WANG X; WANG X T M R I; LI B Y
 PATENT ASSIGNEE: (LIBB-I) LI B; (SHIL-I) SHI L; (SYGN-C) SYNGENTA PARTICIPATIONS AG; (WANG-I) WANG X
 COUNTRY COUNT: 95

PATENT INFO ABBR.:

PATENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC	
WO 2002002805	A2	20020110	(200214)*	EN	76[0]		<--
AU 2001079704	A	20020114	(200237)	EN			<--
US 20030148276	A1	20030807	(200358)	EN			<--
US 20030165923	A1	20030904	(200359)	EN			<--
EP 1356096	A2	20031029	(200379)	EN			<--
AU 2001279704	B2	20051215	(200654)	EN			<--
US 7202022	B2	20070410	(200726)	EN			<--
US 7300751	B2	20071127	(200780)	EN			<--
CA 2413423	C	20110111	(201119)	EN			<--

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE

WO 2002002805 A2	WO 2001-EP7469 20010629
US 20030148276 A1 Provisional	US 2000-215596P 20000630
US 20030165923 A1 Provisional	US 2000-215596P 20000630
US 7202022 B2 Provisional	US 2000-215596P 20000630
US 7300751 B2 Provisional	US 2000-215596P 20000630
AU 2001079704 A	AU 2001-79704 20010629
AU 2001279704 B2	AU 2001-279704 20010629
EP 1356096 A2	EP 2001-957906 20010629
US 20030148276 A1	US 2001-896324 20010629
US 20030165923 A1 CIP of	US 2001-896324 20010629
US 7202022 B2	US 2001-896324 20010629
US 7300751 B2 CIP of	US 2001-896324 20010629
EP 1356096 A2	WO 2001-EP7469 20010629
US 20030165923 A1	US 2002-236363 20020906
US 7300751 B2	US 2002-236363 20020906
CA 2413423 C	CA 2001-2413423 20010629
CA 2413423 C PCT Application	WO 2001-EP7469 20010629

FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 2001079704 A	Based on	WO 2002002805 A
EP 1356096 A2	Based on	WO 2002002805 A
AU 2001279704 B2	Based on	WO 2002002805 A
CA 2413423 C	Based on	WO 2002002805 A

PRIORITY APPLN. INFO: US 2000-215596P 20000630
 US 2001-896324 20010629
 US 2002-236363 20020906

AN 2002-106604 [200214] WPIDS
 AB WO 2002002805 A2 UPAB: 20110321

NOVELTY - A new method (M1) for the simultaneous sequence-specific identification and separation of polynucleotide fragments in a polynucleotide population (P1) comprises using restriction endonucleases that recognize degenerate bases in their recognition or cleavage sequence to produce P1 from transcribed RNA. The fragments are then ligated and amplified.

DETAILED DESCRIPTION - A new method (M1) for the simultaneous sequence-specific identification and separation of polynucleotide fragments in a polynucleotide population (P1) comprises using restriction endonucleases that recognize degenerate bases in their recognition or cleavage sequence to produce P1 from transcribed RNA. The fragments are then ligated and amplified.

In detail, M1 comprises:

- (a) reverse transcribing an RNA population to provide P1;
- (b) digesting P1 with one or more restriction endonucleases (RE1) having a degenerate recognition or cleavage sequence comprising a degenerate base, where the degenerate base is represented by the formula of Nm, where N is the extent of degeneracy, and m is the number of degenerate bases, to produce restriction fragments having Nm different single-stranded overhangs for each restriction endonuclease;
- (c) ligating the restriction fragments having the same overhangs to a series of adapters whose sequences are complementary to the overhangs; and
- (d) amplifying the restriction fragments.

INDEPENDENT CLAIMS are included for the following:

- (1) a method (M2) for detecting polymorphism comprising:
 - (a) reverse transcribing an RNA population to provide a polynucleotide population;
 - (b) digesting the polynucleotide population with one or more RE1;
 - (c) ligating the restriction fragments having the same overhangs to

a series of adapters whose sequences are complementary to the overhangs;

(d) amplifying the restriction fragments;

(e) sequencing the amplified restriction fragments; and

(f) comparing the sequence of the amplified restriction fragments with the sequence of the same polynucleotide from a different source;

(2) a method (M3) for screening for interactions between a preselected protein and polypeptide fragments, comprising culturing the recombinant host cells of (1) under conditions which enable expression of correctly inserted restriction fragments by the host cell, and assaying the interaction of the polypeptide fragments encoded by the restriction fragments with the preselected protein;

(3) a method (M4) for detecting a change in the pattern of RNA expression in a tissue or cell associated with an internal or external factor comprising:

(a) determining the pattern of RNA expression in a first tissue or cell sample not subject to the internal or external change by a method comprising:

(i) reverse transcribing an RNA population to provide the polynucleotide population;

(ii) digesting a double-stranded cDNA library prepared from the RNA isolated from the first sample with one or more RE1, where m is 1-5;

(iii) ligating the restriction fragments produced from (ii) having the same overhangs to a series of adapters whose sequences are complementary to the overhangs;

(iv) amplifying the restriction fragments; and

(v) displaying the pattern of RNA expression in the first sample;

(b) determining and optionally quantifying the pattern of RNA expression in a second tissue or cell sample subject to the physiological or pathological change by performing steps (a)-(d) of M1 with the second sample;

(c) digesting the restriction fragments obtained in step (b) with one or more further restriction endonucleases producing restriction fragments with single-stranded overhangs different from those produced in step (b); and

(d) comparing the first and the second displays to determine the effect of the internal or external factor on the pattern of RNA expression in the tissue;

(4) a method (M5) for diagnosis of a disease based on detecting a change in the pattern of DNA fragments in a disease tissue or cell sample comprising:

(a) determining the pattern of DNA fragments in the sample by:

(i) digesting a DNA isolated from the tissue or cell with one or more restriction RE1;

(ii) ligating the restriction fragments produced from (i) having the same overhangs to a series of adapters whose sequences are complementary to the overhangs;

(iii) amplifying the restriction fragments; and

(iv) displaying the pattern of DNA fragments in the disease tissue or cell sample;

(b) determining the pattern of DNA fragments in a normal tissue or cell corresponding to the disease tissue by performing steps (a)-(d) of M1 with the normal tissue; and

(c) comparing the disease and the normal tissue or cell profile of DNA fragments;

(5) a method (M6) for detecting a change in the pattern of RNA expression in a cell sample in response to an external factor comprising steps (a) to (d) of M4;

(6) a method (M7) for constructing an expressed sequence tag (EST) library comprising:

(a) reverse transcribing an mRNA population isolated from a eukaryotic source to provide a polynucleotide population;

(b) digesting a double-stranded cDNA library prepared from the mRNA

with one or more RE1;

(c) ligating the restriction fragments produced from (b) having the same overhangs to a series of adapters whose sequences are complementary to the overhangs;

(d) amplifying the restriction fragments; and

(e) ligating the amplified restriction fragments into a suitable cloning vector;

(7) an isolated nucleic acid comprising an oligonucleotide selected from one of the 64 nucleotide sequences defined in the specification; and

(8) a recombinant host cell transformed with a vector comprising cloned polymerase chain reaction (PCR) obtained from M1.

USE - The method is useful for the simultaneous sequence-specific identification, separation and quantitative measurement of polynucleotide fragments in a polynucleotide population. The method is applicable in DNA fingerprinting, differential display of mRNA, mutation and polymorphism identification, drug screening, molecular taxonomy, and diagnosis of diseases such as heart disease, lung disease, kidney disease, neurodegenerative disease, liver disease, a disease of the reproductive system, or cancer.

The isolated nucleic acid molecule comprising an oligonucleotide selected from one of the 64 nucleotide sequences defined in the specification can be used in any of the methods described above (claimed).

ADVANTAGE - The method is rapid, quantitative and it determines the gene expression level without the requirement of sequence information.

L4 ANSWER 5 OF 6 WPIDS COPYRIGHT 2012 THOMSON REUTERS on STN
ACCESSION NUMBER: 2002-048724 [200206] WPIDS
CROSS REFERENCE: 2001-040475; 2001-366407; 2002-617299
DOC. NO. CPI: C2002-013582 [200206]
TITLE: Isolating DNA containing fragments nicked by Escherichia coli methyl-directed mismatch repair system involves using a modified rolling circle amplification procedure which employs DNA polymerase III
DERWENT CLASS: B04; D16
INVENTOR: LASKEN R; WEISSMAN S
PATENT ASSIGNEE: (LASK-I) LASKEN R; (WEIS-I) WEISSMAN S
COUNTRY COUNT: 1

PATENT INFO ABBR.:

PATENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC
US 20010039039	A1	20011108	(200206)*	EN	19[5]	<--

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
US 20010039039	A1 Provisional	US 1998-100996P	19980918
US 20010039039	A1 Div Ex	US 1999-398216	19990917
US 20010039039	A1	US 2001-820356	20010329

FILING DETAILS:

PATENT NO	KIND	PATENT NO
US 20010039039	A1 Div ex	US 6235502 B

PRIORITY APPLN. INFO: US 2001-820356 20010329
US 1998-100996P 19980918
US 1999-398216 19990917

AN 2002-048724 [200206] WPIDS

CR 2001-040475; 2001-366407; 2002-617299
AB US 20010039039 A1 UPAB: 20060118

NOVELTY - Mismatched DNA containing fragments nicked by MutSLH from two DNA samples are isolated by a modified rolling circle amplification (RCA) method. DNA sample is digested with restriction enzyme, the fragments are ligated to Y-shaped adapters forming adapter-fragment constructs which are nicked by MutSLH proteins and nicked fragments are treated with DNA polymerase(s) which elongates the 3'-OH end.

DETAILED DESCRIPTION - Isolating (M1) mismatched DNA containing fragments nicked by MutSLH (a combination of the three proteins of the Escherichia coli methyl-directed mismatch repair system, collectively called MutSLH) from two DNA samples, comprises:

- (a) digesting a first DNA sample with a restriction enzyme to obtain DNA fragments;
- (b) ligating Y-shaped adapters to the fragments to obtain fragment-plus-adapter constructs;
- (c) repeating the above steps with a second sample;
- (d) methylating the products formed from (c) of the first sample but not the second sample;
- (e) mixing the methylated and unmethylated products obtained from the samples, denaturing and reannealing to form hemimethylated heterohybrids;
- (f) treating the fragments produced in step (e) with a MutSLH preparation to nick DNA containing mismatches and form a 3'-OH end; and
- (g) elongating the 3'-OH end with one or more DNA polymerases.

The method can also be carried out by ligating adapters to DNA fragments to obtain fragment-adapter constructs that have a single-stranded (ss) overhang on the 5'-ends and recessed 3'-ends, where the adapters are blocked with a dideoxynucleotide or other modification that prevents the elongation by a DNA polymerase, carrying out (c)-(e) as above, blocking pre-existing nicks on the fragments with dideoxynucleoside triphosphates or their analogs, treating the fragments with MutSLH preparation to form 3'-OH ends and elongating the ends with a DNA polymerase having strand displacement or nick translation capacity.

INDEPENDENT CLAIMS are also included for the following:

(1) amplifying (M2) DNA using RCA in a manner that results in approximately equimolar amplification of the DNA, by amplifying a circle DNA molecule which comprises a stem-and-loop, or a circle DNA molecule containing a universal base, an abasic residue, or other residue that alters the rate of circle replication, where elongation of the circle by a replicating polymerase carrying out the amplification is inhibited;

(2) a method (M3) for using RCA to obtain approximately equimolar amplification of DNA fragment mixtures by introducing at least one pause site in a DNA circle, comprises: (a) placing within the circle a stretch of one or more abasic sites, a region of high secondary structure and a ligand binding site, and then carrying out RCA under conditions such that the pause site slows the DNA polymerase replicating the circle and tends to equalize the number of times the polymerase copies circles of different length; (b) circularizing dsDNA fragments around an adapter such that one strand of the adapter has an internal pause and the outer strand is blocked from ligation at one or both ends; or (c) using a splint that has an internal double stranded (ds) segment containing a pause site on one DNA strand, and which has single stranded extensions at both ends complementary to the ends of the single-stranded fragment to be amplified;

(3) a method (M4) in which fragments of DNA may be circularized and amplified, by cutting DNA with a restriction enzyme that produces a single fragment containing all the markers of the region having pause to be determined, or using the RecA-assisted restriction endonuclease (RARE) method or its variants to produce such a fragment, ligating the cut DNA around an adapter and amplifying the fragment using RCA to obtain a ssDNA from the desired fragment;

(4) amplifying (M5) segments of a DNA target by RCA, by annealing a

padlock DNA that creates a gap comprising the sequence to be amplified, extending the 3'-OH of the padlock DNA with a DNA polymerase which generates a nick, ligating the nick to form a circle DNA, annealing an RCA primer to the circle DNA and carrying out RCA with DNA polymerase III;

(5) a method (M6) in which large fragments of DNA may be circularized and amplified, by providing a DNA-adapter construct which has a DNA fragment ligated around an adapter that has an internal gap and 5' region of non-complementarity and amplifying the DNA-adapter construct using DNA pol III in a RCA;

(6) ligating (M7) adapters onto ends of DNA fragments for use in RCA of the DNA, by ligating restriction endonuclease fragments with an oligonucleotide adapter that forms a hairpin structure with itself such that 3' and 5' ends of the adapter are annealed together and form a proper overhang or blunt end for ligation to the restriction endonuclease fragments, where a circular construct is formed, annealing an oligonucleotide primer to the single-stranded loop portion of the adapters and extending the oligonucleotide primer with a DNA polymerase in RCA reaction; and

(7) carrying out RCA reaction, where at least two different DNA polymerases are employed, one or more of which contains a 3' to 5' exonuclease activity capable of correcting nucleotide misincorporations.

USE - (M1) is useful for isolating mismatched DNA containing fragments nicked by MutSLH from two DNA samples. The procedure allows for the MutSLH nicking of DNA fragments containing mismatched bases to analyze polymorphism between DNA samples. The methods are useful in phase determination, polymorphism analyses, mismatch scanning procedures and cloning procedures, diagnostics (claimed), genotyping, genomic mapping, DNA sequencing and synthesis of DNA probes.

ADVANTAGE - DNA polymerase III holoenzyme provides a superior rate of DNA synthesis and also high processivity which allows rapid replication through regions of high GC content, hair-pin structures, and other regions of secondary structure and regions that normally slow replication due to local sequence context effects.

DESCRIPTION OF DRAWINGS - The figure shows the fragment plus adapter constructs obtained by ligating adapters to DNA fragments, in the amplification of specific DNA sequences by rolling circle amplification.

L4 ANSWER 6 OF 6 WPIDS COPYRIGHT 2012 THOMSON REUTERS on STN
ACCESSION NUMBER: 1997-132667 [199712] WPIDS
DOC. NO. CPI: C1997-042908 [199712]
TITLE: Production, and opt. amplification, of hairpin nucleic acid
using a single primer - that is extended to form a
template for synthesis of extension product, useful e.g.
in gene walking and sequencing
DERWENT CLASS: B04; D16
INVENTOR: DEWHIRST F E
PATENT ASSIGNEE: (FORS-N) FORSYTH DENTAL INFIRMARY FOR CHILDREN
COUNTRY COUNT: 19

PATENT INFO ABBR.:

PATENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC
WO 9704131	A1	19970206	(199712)*	EN	66[17]	<--

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9704131	A1	WO 1996-US11948	19960718

PRIORITY APPLN. INFO: US 1995-505094 19950721

AN 1997-132667 [199712] WPIDS

AB WO 1997004131 A1 UPAB: 20060112

Production of at least one copy of a hairpin polynucleotide (I) comprises:
(a) combining (I) with a template-dependent polynucleotide polymerase (II); nucleoside triphosphates (NTPs) and excess of one primer (P) that hybridises to a site in the 3' part of (I) and can produce an extension product (EP); (b) dissociating the 2 complementary segments of (I) and (c) allowing hybridisation of P and thus its extension to produce a complementary copy of (I) asan EP.Opt. the process is repeated for amplification.

Also new are kits for the process containing hairpin DNA (the adapter) with a phosphorylated 5' end and ligatable to a blunt or restriction end of double stranded (ds) DNA, plus a ligase, the kits also may contain a polymerase and restriction enzymes.

USE - The method is specified for (1) detecting (I) (from the amplification product) and (2) sequencing ds DNA after its conversion to (I). Typical applications are in gene or chromosome walking; amplification and sequencing of cDNA and detecting even minute amounts of DNA.

ADVANTAGE - Since only a single primer is required, the method allows amplification of flanking regions.

=> log

ALL L# QUERIES AND ANSWER SETS ARE DELETED AT LOGOFF

LOGOFF? (Y)/N/HOLD:hold

COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	161.58	161.82
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-0.87	-0.87

SESSION WILL BE HELD FOR 120 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 16:40:33 ON 04 JAN 2012